INTERNATIONAL SEARCH REPORT

International application No.

PCT/US04/33530

IPC(7)	SSIFICATION OF SUBJECT MATTER A61K 31/535, 31/40					
US CL	: 514/236.2, 411		ifi tro			
	International Patent Classification (IPC) or 6 both na DS SEARCHED	itional cla	ssification and IPC			
						
Minimum documentation searched (classification system followed by classification symbols) U.S 514/236 2, 411						
Documentation searched other than minimum documentation to the extent that such documents are included m the fields searched						
Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) Please See Continuation Sheet						
	UMENTS CONSIDERED TO BE RELEVANT					
Category *	Citation of document, with indication, where a	ppropriate	e, of the relevant passages	Relevant to claim No		
A		Patabase MEDLINE m STN (Columbus, OH, USA), AN 2001045397, KRUM H et al, 1-41, 55-67 Baseline predictors of tolerability to carvedilol in patients with chronic heart failure, Heart, lecember 2000, vol. 84, no 6, pages 615-9, abstract				
Y	Database CAPLUS in STN (Columbus, OH, USA) DN 13 1:252307, NAGARAJA S et al, Treatment with inverse agonists enhances baseline atrial contractility in transgenic mice with chronic beta2-adrenoceptor activation, Br J Pharmacol, 1999, vol.127, no.5, pages 1099-1104, abstract.					
А	Database MEDLINE in STN (Columbus, OH, USA) AN 84081647, MARKOWITZ S et al, "Timolol: a 4-year follow-up study, ¹ Can J Ophthalmol, 18 October 1983, vol, 18, no. 6, pages 278-80, abstract					
A	A Database MEDLINE (Columbus, OH, USA), AN 2002624573, SEIFERT R et al, 'Consitutive activity of G-protein-coupled receptors: cause of disease and common property of wild-type receptors,' Naunyn-Schmiedeberg's archives of pharmacol, November 2002, vol. 366, no. 5, pages 381-416, abstract.					
Further	documents are listed in the continuation of Box C.		See patent family annex.			
Special categanes of cited documents "A" document definingthe general state of the art which is not considered to be of particular relevance		T	later document published after the inten and not in conflict with the application be principle or theory underlying the inven-	out cited to understand the		
"E" earlier ap	plication or patent published on or after the international filing date	"X" document of particular relevance, the claimed invention cannot be considered novel or cannot be considered to involve an inventive ste when the document is taken alone		aimed invention cannot be ed to involve an inventive step		
"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)		"Y" document of particular relevance, the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious				
"O" document referring to an oral disclosure, use, exhibition or other means "P" document published pnor to the international filing date but later than the		*&. <i>*</i>	to a person skilled in the art document member of the same patent fa	-		
priority date claimed						
Date of the actual completion of the international search		Date of mailing of the international searcf Lreport 0 4 NUV ZUUD				
28 September 2005 (28.09.2005) Name and mailing address of the ISA/US		Authori	zed officer _ c			
Mail StopPCT, Attn ISA/US Commissioner of Patents		Rebecc		ford		
P O Box 1450			one No. 703-308-1235	£91		
	Facsimile No (703) 305-3230 Form PCT/ISA/210 (second sheet) (July 1998)					
C 1/13/A	and (second sileet) (July 1770)		•	7		

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Categoiy *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.	
Ÿ	Database BIOTECHDS in STN (Columbus, OH, USA) AN 2003-03745, OAKLEY R et al, 'Modified B-protein coupled receptor useful for identifying an agonist, inverse agonist or antagonist of the receptor, comprises a carboxyl terminal having one or more clusters of phosphorylation, vector-mediated gene transfer and expression in host cell for recombinant protein production, drug screening and gene therapy,' US 2002106739, 05 November 2001, abstract.	42-54	
Α	Database CAPLUS in STN (Columbus, OH, USA) DN138:84549, MARTINEZ R et al, 'Protein and cDNA sequences of human olfactory G protein-copied receptors, their diagnostic and therapeutic uses thereof,' WO 2003000735, 03 January 2003, abstract.	1-67	
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INTERNATIONAL SEARCH REPORT

International application No.
PCT/US04/33530

Box I Observations where certain claims were found unsearchable (Continuation of Item 1 of first sheet)			
This international report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:			
Claim Nos.: because they relate to subject matter not required to be searched by this Authority, namely:			
Claim Nos.: because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:			
3. Claim Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).			
Box π Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)			
This International Searching Authority found multiple inventions in this international application, as follows: Please See Continuation Sheet			
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. M			
1. As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.			
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.			
3. As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:			
4. No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:			
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.			

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INTERNATIONAL SEARCH REPORT	PCT/US04/33530
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BOX π . OBSERVATIONS WHERE UNITY OF INVENTION IS LAC Group I, claim(s) 1-41, 55-67, drawn to a method of use, holders and compositions.	CKING
Group H, claim(s) 42-51, drawn to a first method of screening in which the basal le measured.	evel of activity of the compound that is screened is
Group III, claim(s) 52-54, drawn to a second method of screening in which receptor is screened is measured.	population or receptor density of the compound that
The inventions listed as Groups I-HI do not relate to a single general inventive conc 13.2, they lack the same or corresponding special technical features for the followir while Groups II and III are drawn to methods of detecting activity of compounds. F require the elements of the screening method of Group HI.	ng reasons: Group I is to a method of treating a disease
Continuation of B. FIELDS SEARCHED Item 3: REGISTRY, CAPLUS, MEDLINE, DRUGU, BIOSIS, IPA, TOXCENTER, EMBA	SE. JICST-EPLUS. PASCAL, BIOTECHNO
ESBIOBASE, CONFSCI, BIOTECHDS, DISSABS, TOXCENTER, WPIDS, SCISI	